

SEQUENCE LISTING

<110> Blenis, John
Murphy, Leon O.

<120> Inhibitors of the Map Kinase Pathway

<130> 00742/074002

<140> US 10/563,345

<141> 2006-01-03

<150> PCT/US04/21514

<151> 2004-07-02

<150> US 60/484,761

<151> 2003-07-03

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Gly Pro Met Val Thr Glu Leu Glu Pro Leu Cys Thr Pro Val Val Thr

1

5

10

15

Cys Thr Pro Ser Cys Thr Thr Tyr Thr Ser Ser Phe Val Phe Thr Tyr

20

25

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Pro Glu Glu Ala Asp Ser

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Gly Pro Val Leu Glu Pro Glu Ala Leu His Thr Pro Thr Leu Met Thr

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Ser	Thr	Pro	Glu	Pro											
		35													

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Ser Thr Pro Ala Ile Thr Pro Gly Thr Ser Asn Leu Val Phe Thr Tyr
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Pro Ser Val Leu Glu Gln
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His Ser Glu Val Gln Val Leu Gly Asp Pro Phe Pro Val Val Ser Pro
1 5 10 15
Ser Tyr Thr Ser Ser Phe Val Leu Thr Cys Pro Glu Val Ser Ala Phe
20 25 30

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Leu Leu Ala Ser Pro Asp Leu Gly Leu Leu Lys Leu Ala Ser Pro Glu
1 5 10 15
Leu Glu Arg Leu Ile Ile Gln Ser Asn Gly Leu Val Thr Thr Thr Pro
20 25 30
Thr Ser Thr Gln Phe Leu Tyr Pro Lys Val
35 40

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Gly Gln Gly Ser Asp Thr Gly Ala Ser Leu Lys Leu Ala Ser Ser Glu
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Leu Glu Arg Leu Ile Val Pro Asn Ser Asn Gly Val Ile Thr Thr Thr
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Pro Thr Pro Pro Gly Gln Tyr Phe Tyr Pro Arg Gly
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 Leu Glu Arg Leu Ile Ile Gln Ser Ser Asn Gly His Ile Thr Thr Thr
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 Pro Thr Pro Thr Gln Phe Leu Cys Pro Lys Asn
 35 40

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<400> 13
 Leu Pro Thr Pro Pro Leu Ser Pro Ser Arg Arg Ser Gly Leu Cys Ser
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 Pro Ser Tyr Val
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 Leu Thr Ala Ala Ala Ser Glu Cys Ile Asp Pro Ser Val Val Phe Pro
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 Tyr Pro Leu Asn Asp
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 Ala Gln Ser Pro Gly Ala Gly Ala Ala Ser Pro Ala Gly Arg Gly His
 1 5 10 15
 Gly Gly Ala Ala Gly Ala
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<400> 16

Ala	His	Pro	Ala	Ala	Glu	Cys	Val	Asp	Pro	Ala	Val	Val	Phe	Pro	Phe
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Pro	Val	Asn	Lys												
			20												

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Gln	Ser	Pro	Pro	Leu	Ser	Cys	Ala	Val	Pro	Ser	Asn	Asp	Ser	Ser	Pro
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Ile	Tyr	Ser	Ala	Ala	Pro	Thr	Phe	Pro	Thr	Pro	Asn	Thr	Asp		
			20					25					30		

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Pro	Met	Ile	Pro	Asp	Tyr	Leu	Phe	Pro	Gln	Gln
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<400> 19

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Pro	Ala	Ala	Phe	Pro	Ala	Pro	Leu	Val	Thr	Pro	Met	Val	Ala	Leu	Pro
			20					25					30		
Asn	Tyr	Leu	Phe	Pro	Thr	Pro	Ser	Ser	Tyr						
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Gln Phe Pro

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<400> 22
Arg Arg Pro Arg Ser Pro Ala Lys Leu Ser Ala Gln Ala Pro Ser Ala
1 5 10 15
Gln Ala Pro

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Arg Arg Pro Arg Ser Pro Ala Lys Leu Ser Phe Thr Tyr Pro Ser Phe
1 5 10 15
Thr Tyr Pro

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Arg Arg Pro Arg Ser Pro Ala Lys Leu Ser Ala Thr Tyr Pro Ser Ala
1 5 10 15
Thr Tyr Pro

<210> 25
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Cys Thr Pro Ser Cys Thr Thr Tyr Thr Ser Ser Phe Val Phe Thr Tyr
20 25 30
Pro Glu Ala Asp Ser Phe Pro Ser Cys Ala Ala Ala His Arg Lys Gly

35 40 45
 Ser Ser Ser Asn Glu Pro Ser Ser Asp Ser Leu Ser Ser Pro Thr Leu
 50 55 60
 Leu Ala Leu
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<400> 26
 Gly Glu Glu Pro Leu His Thr Pro Ile Val Val Thr Ser Thr Pro Ala
 1 5 10 15
 Val Thr Pro Gly Thr Ser Asn Leu Val Phe Thr Tyr Pro Ser Val Leu
 20 25 30
 Glu Gln Glu Ser Pro Ala Ser Pro Ser Glu Ser Cys Ser Lys Ala His
 35 40 45
 Arg Arg Ser Ser Ser Ser Gly Asp Gln Ser Ser Asp Ser Leu Asn Ser
 50 55 60
 Pro Thr Leu Leu Ala Leu
 65 70

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<400> 27
 Gly Pro Val Leu Glu Pro Glu Ala Leu His Thr Pro Thr Leu Met Thr
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 Thr Pro Ser Leu Thr Pro Phe Thr Pro Ser Leu Val Phe Thr Tyr Pro
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 Ser Thr Pro Glu Pro Cys Ser Ser Thr His Arg Lys Ser Ser Ser Ser
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 Ser Gly Asp Pro Ser Ser Asp Pro Leu Gly Ser Pro Thr Leu Leu Ala
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 Leu
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Thr Gln Phe Trp Phe Trp Phe Pro Leu Ser His Phe Leu Ser Leu
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<400> 32
Lys Pro Ser Thr Phe Ala Tyr Pro Ala Pro Leu Glu Val Pro Lys
1 5 10 15

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Glu	Gly	Lys	Leu	Trp	Glu	Phe	Pro	Ile	Asn	Asn	Glu	Ala	Gly	Phe
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Ile	Lys	Leu	Glu	Tyr	Asp	Phe	Pro	Glu	Lys	Phe	Phe	Pro	Lys	Ala
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Val	Gln	Lys	Arg	Phe	Gly	Phe	Pro	Glu	Gly	Ser	Val	Glu	Leu	Tyr
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Asp	Leu	Ala	Phe	Phe	Arg	Phe	Pro	Arg	Asp	Pro	Ala	Arg	Cys	Gln
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<400> 37

Pro	Thr	Val	His	Tyr	Glu	Phe	Pro	Asn	Gly	Tyr	Asn	Cys	Asp	Phe
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Asp	Leu	Pro	Lys	Tyr	Leu	Phe	Pro	Glu	Asp	Pro	Ser	Tyr	Ser	Ser
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Asp Leu Pro Ser Tyr Leu Phe Pro Glu Asp Pro Ser Tyr Asp Ala
1 5 10 15

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1 5 10 15

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Gly Leu Thr Pro Phe Ser Phe Pro Gln Val Thr Asn Ser Ser Val
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Glu Pro Cys Pro Phe Ser Phe Pro Met Pro Tyr Pro Asn Tyr Arg

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 Ala Ser Thr Gly Tyr Pro Phe Pro Met Val Phe Ser Ser Cys Ser
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 Pro Asp Asn Trp Phe Cys Tyr Pro Cys Ile Pro Asn His Glu Leu
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<400> 48
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Arg Pro Ser Leu Phe Val Tyr Pro Glu Glu Ser Leu Val Ile Gly
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Gly Ile Thr Val Phe Tyr Phe Pro Arg Cys Gln Gly Phe Thr Ser
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Ser Ser Gly Ile Phe Gln Phe Pro Leu Asn Leu Cys Thr Lys Thr
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Asp Thr Glu Gly Phe Lys Tyr Pro Leu Gly His Glu Val Asn His
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Gln Asp Phe Ser Tyr Phe Phe Pro Asp Asp Pro Pro Thr Phe Ile
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Glu Glu Gly Pro Phe Thr Phe Pro Asn Gly Glu Ala Val Glu His
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Glu Ala Leu Glu Phe Tyr Tyr Pro Glu His Phe Thr Leu Leu Thr
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Glu Ser Leu Glu Phe Tyr Tyr Pro Glu Leu Tyr Lys Leu Val Thr
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Leu Pro Arg Asn Phe His Tyr Pro Pro Asp Gly Tyr Ser Arg His
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Pro Glu Gly Ser Phe Gln Tyr Pro Ala Ser Tyr His Ser Asn Gln
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Ser Ser Ala Ala Phe Gly Phe Pro Arg Gly Ala Gly Pro Ala Gln
1 5 10 15

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1 5 10 15

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Ser Pro Ala Pro Phe Pro Phe Pro Glu Ala Pro Gly Ser Leu Pro
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Arg Ser Leu Ile Tyr Lys Phe Pro Ser Thr Phe Tyr Glu Gly Arg
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Ser Tyr Ala Lys Phe Leu Tyr Pro Thr Asn Ala Leu Val Thr His
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Glu Tyr Phe Asp Tyr Ile Phe Pro Glu Asp Ala Ala Asn Gln Pro
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Thr Glu Asp Asp Phe Glu Phe Pro Phe Ala Lys Thr Asn Leu Ser
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Pro Gln Ala Pro Phe Gly Tyr Pro Gly Asp Gly Met Gln Gln Pro
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 Thr Arg Phe Ser Tyr Ala Phe Pro Lys Glu Phe Pro Tyr Arg Met
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 Ser Ser Leu Asn Phe Ser Phe Pro Ser Leu Pro Thr Met Gly Gln
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Met	Thr	Tyr	Leu	Phe	Asp	Phe	Pro	His	Gly	Pro	Lys	Pro	Gly	Ser
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Met	Glu	Arg	Val	Phe	Gly	Phe	Pro	Val	His	Tyr	Thr	Asp	Val	Ser
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Leu	Glu	Arg	Ile	Phe	Gly	Phe	Pro	Val	His	Tyr	Thr	Asp	Val	Ser
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His Val Asn Ala Phe Gly Phe Pro Pro Thr Glu Pro Ser Ser Thr
1 5 10 15

<210> 98
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Asn Lys Leu Tyr Tyr Asn Phe Pro Trp Gly Lys Glu Leu Ile Glu
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His Arg Ile Gln Phe Lys Tyr Pro Gly Pro Glu Asp Asp Ala Ala
1 5 10 15

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 Gly Ser His Leu Phe Gly Phe Pro Pro Thr Pro Pro Lys Glu Val
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Val	Tyr	Asn	Ser	Phe	Gly	Phe	Pro	Gly	Gln	Ala	Ala	Arg	Tyr	Pro
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Glu	Pro	Glu	Asp	Phe	Ala	Phe	Pro	Ser	Thr	Ala	Pro	Ser	Pro	Gln
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Ala	Pro	Gln	Pro	Tyr	Thr	Phe	Pro	Ala	Ser	Leu	Ser	Thr	Ile	Asn
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Thr	Ala	Leu	Gln	Phe	Thr	Tyr	Pro	Leu	Phe	Thr	Thr	Asn	Ala	Cys
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Pro	Pro	Glu	Asn	Tyr	Asp	Phe	Pro	Val	Val	Ile	Val	Lys	Gln	Glu
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<400> 283

Gly	Gly	Ala	Ala	Phe	Ile	Phe	Pro	Asn	Thr	Ser	Val	Tyr	Pro	Glu
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Asn	Val	Pro	Asp	Phe	Asp	Phe	Pro	Pro	Glu	Phe	Tyr	Glu	His	Ala
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Phe	Pro	Val	Arg	Phe	Pro	Tyr	Pro	Cys	Thr	Gln	Thr	Glu	Leu	Ala
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Gly	Leu	Glu	Thr	Tyr	Gly	Phe	Pro	Ser	Val	Thr	Leu	Ile	Ser	Cys
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<400> 318

Ser	Thr	Asn	Pro	Phe	Asp	Phe	Pro	Phe	Val	Ser	Gln	Gly	Glu	Val
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<213> Homo sapiens

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Phe	Pro	Gln	Gly	Tyr	Gln	Phe	Pro	Ser	Met	Glu	Gln	Leu	Ala	Ala
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Pro	Met	Asp	Arg	Phe	Ser	Phe	Pro	Asp	Gly	Lys	Glu	Ser	Asp	Thr
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Asp	Val	Ala	Asp	Tyr	Val	Phe	Pro	Ala	Gln	Gly	Asp	Asn	Ser	Phe
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Asp	Thr	Ala	Asp	Tyr	Thr	Phe	Pro	Leu	Gln	Gly	Asn	Ser	Phe	Phe
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Val	Phe	Val	Gly	Phe	Leu	Phe	Pro	Trp	Ala	Leu	Met	Leu	Leu	Ser
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